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Claims:

- A method for measuring a blood oxygen level-dependent magnetic resonance imaging signal, comprising
- a) administering an effective amount of an allosteric effector compound capable of decreasing hemoglobin binding affinity for oxygen; and
- b) performing a blood oxygen level-dependent magnetic resonance imaging scan,
 whereby said blood oxygen level-dependent magnetic resonance imaging signal is measured.
- The method of Claim 1, wherein the allosteric effector compound is selected from the group consisting of:

a compound having the formula:

where $R_{1.5}$ may be hydrogen, halogen, or a substituted or unsubstituted $C_{1.3}$ alkyl group and may be the same or different,

 R_{6-7} may each be hydrogen or methyl and may be the same or different, and

 R_8 may be hydrogen, a substituted or unsubstituted $C_{1\text{-}3}$ alkyl group, or a salt cation, and X and Z are CH₂, NH, or O;

a compound having the formula:

where X and Z may each be CH_2 , CO, NH or O, and Y may be CO or NH, which the caveat that X, Y, and Z must all be different from each other, and

 $R_{2\text{-}6}$ can be the hydrogen, halogen, substituted or unsubstituted $C_{1\text{-}3}$ alkyl groups, and may be the same or different,

 $R_{7.8}$ can be hydrogens, methyls, ethyls, or alkyl groups in a ring connecting the two, and R_9 can be a hydrogen, lower alkyl, or salt cation;

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$$R_{4}$$

$$R_{5}$$

$$R_{6}$$

$$R_{6}$$

$$R_{1} = O - C - COOR_{9}$$

$$R_{8}$$

where $R_{3\cdot6}$ can be the hydrogen, halogen, substituted or unsubstituted $C_{1\cdot3}$ alkyl group, or a $C_{1\cdot3}$ ether or ester, and these moieties may be the same or different, or alkyl moieties of an aromatic or aliphatic ring incorporating two of the $R_{3\cdot6}$,

R1 can be connected to any position on the phenyl ring, and

sites $R_{7.8}$ can be hydrogen, halogen, methyl, ethyl, and these moieties may be the same or different, or alkyl groups in a ring connecting the two, and

 R_9 can be a hydrogen, halogen, C_{1-3} lower alkyl, or salt cation; a compound having the formula:

$$R_2 - X$$

$$Z$$

$$R_1 = O - C - COOR_9$$

$$R_8$$

where R1 can be connected to any position on the phenyl ring, and

sites $R_{7.8}$ can be hydrogen, halogen, methyl, ethyl, and these moieties may be the same or different, or alkyl groups in a ring connecting the two, and

 R_2 is defined as a substituted or unsubstituted aromatic compound, a substituted or unsubstituted alkyl ring compound, or a substituted or unsubstituted phthalimide compound,

X is a carboxyl,

Y is a nitrogen,

and R_2 completes the phthalimide compound by being bonded to both X and Y; and where X, Y, and Z, may either be CH_2 , NH, O, or N, with the caveat that each are different from the other:

$$R_3$$
 R_4
 R_6
 R_7
 R_8
 R_7
 R_8
 R_7
 R_8

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where R2, R3, R4, R5, and R6 may be hydrogen, halogen, or alkyl groups and may be the same or different.

 R_7 and R_8 may be hydrogen or methyl groups and may be the same or different, and where the R_9 moiety is hydrogen or a salt cation;

a compound having the formula:

where R_2 is a substituted or unsubstituted aromatic compound, or a substituted or unsubstituted alkyl ring compound, or a substituted or unsubstituted phthalimide compound that incorporates X and Y,

X is a carbonyl,

Y is a nitrogen, and

 R_2 completes the phthalimide compound by being bonded to both X and Y, and where X, Y, and Z are CH_2 , NH, S, SO_2 , CO, O or N with the caveat that X, Y, and Z are each different from one another, and

where R1 can be connected to any position on the phenyl ring, and

R₃ and R₄ are hydrogen, halogen, methyl, ethyl, propyl, isopropyl, neopentyl, butyl, or substituted or unsubstituted aryl groups and these moieties may be the same or different, or alkyl moieties as part of an aliphatic ring connecting R₃ and R₄, and

Rs is a hydrogen, halogen, C1-3 lower alkyl, or a salt cation;

a compound having the formula:

$$R_8$$
 R_7 R_1 R_2 R_3 R_6 R_6 R_6 R_4

where A is a chemical bridge which includes two to four chemical moieties bonded together,

the chemical moieties in A are selected from the group consisting of CO, O, S, SO₂, NH, NR₉ where R₉ is a C₁₋₆ alkyl group, CH₂, CH, and C, with the proviso that, except in the case where A contains two identical CH and C moieties positioned adjacent one another to form an alkene or alkyne, the chemical moieties in A are each different from one another, and

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at least one of R₁₋₅ is substituted with a compound having the chemical formula:

$$O-(CH_2)_n$$
 $C-COOR_{12}$
 R_{11}

where n is zero to five.

where R_{10} and R_{11} are selected from the group consisting of hydrogen, halogen, $C_{1\cdot 12}$ alkyl groups, carboxylic acids and esters, aromatic or heteroatomic groups, and these moieties may be the same or different, or alkyl moieties of part of an aliphatic ring connecting R_{10} and R_{11} , and where R_{12} is a hydrogen, halogen, salt cation, metal, or $C_{1\cdot 6}$ alkyl group, and wherein a remainder of the $R_{1\cdot 5}$ moieties and the $R_{6\cdot 8}$ moieties are selected from the group consisting of hydrogen, halogen, $C_{1\cdot 6}$ alkyl groups, $C_{1\cdot 6}$ ether or esters, aromatics and heteroaromatics, and alkyl moieties of an aliphatic ring connecting two sites on a phenyl group;

a compound having the formula:

$$R_1$$
— A — R_2

where R_1 and R_2 each are a substituted or unsubstituted aromatic or heteroaromatic compounds, or a substituted or unsubstituted alkyl or heteroalkyl ring compound, or a substituted or unsubstituted phthalimide compound, and

where R1 and R2 may be the same or different,

where A is a chemical bridge which includes three chemical moieties bonded together between R_1 and R_2 ,

wherein the chemical moieties in A are selected from the group consisting of CO, O, S, SO₂, NH, NR₃ where R_3 is C_{1-6} alkyl group, NR₄ where R_4 includes two carbonyls as part of a phthalimide compound formed with R_1 or R_2 , CH₂, CH, and C, and

where at least one of \mathbf{R}_1 and \mathbf{R}_2 is substituted with a compounds having the chemical formula:

$$O-(CH_2)_n$$
 $-C-COOR_7$
 R_0

where n is zero to five, where R_5 and R_6 are selected from the group consisting of hydrogen, halogen, substituted or unsubstituted C_{1-12} alkyl groups, carboxylic acid and ester groups, substituted or unsubstituted aromatic or heteroaromatic groups, and these moieties may be the same or different, or alkyl moieties of part of an aliphatic ring connecting R_5 and R_6 , and

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where R_7 is a hydrogen, halogen, salt cation, metal, or substituted or unsubstituted $C_{1.6}$ alkyl group;

a compound having the formula:

$$R_1 - A - R_2$$

where R_1 and R_2 each are a substituted or unsubstituted aromatic or heteroaromatic compound, or substituted or unsubstituted alkyl or heteroalkyl ring compound, or a substituted or unsubstituted phthalimide compound, and

where R1 and R2 may be the same or different,

where A is a chemical bridge which includes two to four chemical moieties bonded together between R₁ and R₂,

wherein said chemical moieties in A are selected from the group consisting of CO, O, S, SO_2 , NH, NR_3 where R_3 is a C_{1-6} alkyl group, NR_4 where R_4 includes two carbonyls as part of a phthalimide compound formed with R_1 or R_2 , CH₂, CH, and C, with the caveat that, except in the case where A contains two identical CH and C moieties positioned adjacent one another to form an alkene or alkyne, the chemical moieties in A are each different from one another, and

wherein at least one of R_1 or R_2 is substituted with a compound having the chemical formula:

$$O-(CH2)n-C-COOR7$$

where n is zero to five,

where R_5 and R_6 are selected from the group consisting of hydrogen, halogen, substituted or unsubstituted C_{1-12} alkyl groups, carboxylic acid and ester, substituted or unsubstituted aromatic or heteroaromatic groups, and these moieties may be the same or different, or alkyl moieties of part of an aliphatic ring connecting R_5 and R_6 , and

where R_7 is a hydrogen, halogen, salt cation, metal, or substituted or unsubstituted $C_{1.6}$ alkyl group; and

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where R_1 is selected from the group consisting of optionally substituted phenyl, adamantyl, napthyl, and indanyl, $R_{2\cdot 3}$ are alkyl moieties of a $C_{3\cdot 6}$ alkyl ring connecting R_2 and R_3 , and R_4 is a hydrogen, a monovalent salt cation, or a $C_{1\cdot 3}$ lower alkyl.

- The method of Claim 2, wherein the allosteric effector compound is administered at a dose of 100-300 mg/kg.
- The method of Claim 1, wherein the allosteric effector compound is 2-[4-(((3,5-dimethylanilino)carbonyl)methyl)phenoxy]-2-methylpropionic acid, or a physiologically acceptable salt thereof.
- The method of Claim 4, wherein the allosteric effector compound is administered at a dose of 100-300 mg/kg.
- The method of Claim 4, wherein the allosteric effector compound is administered at a dose of 200 mg/kg.
- A method of increasing the sensitivity of cells to the cytotoxic effects of ionizing radiation comprising:
 - a) contacting the cells with an amount of a compound effective to oxygenate the cells;
 - b) measuring the oxygenation of the cells; and
 - c) administering an effective cytotoxic dose of ionizing radiation to the cells.
- The method of Claim 7, wherein the allosteric effector compound is selected from the group consisting of:

a compound having the formula:

$$R_4$$
 R_5
 R_6
 R_6
 R_7
 R_7
 R_6
 R_6

where $R_{1.5}$ may be hydrogen, halogen, or a substituted or unsubstituted $C_{1.3}$ alkyl group and may be the same or different,

R₆₋₇ may each be hydrogen or methyl and may be the same or different, and

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 R_8 may be hydrogen, a substituted or unsubstituted $C_{1\text{-}3}$ alkyl group, or a salt cation, and X and Z are CH_2 , NH, or O;

a compound having the formula:

$$R_3$$
 R_2 R_4 $X-Y-Z$ $O-C-COOR_9$ R_8

where X and Z may each be CH_2 , CO, NH or O, and Y may be CO or NH, which the caveat that X, Y, and Z must all be different from each other, and

 $R_{2.6}$ can be the hydrogen, halogen, substituted or unsubstituted $C_{1.3}$ alkyl groups, and may be the same or different,

 $R_{7.8}$ can be hydrogens, methyls, ethyls, or alkyl groups in a ring connecting the two, and R_9 can be a hydrogen, lower alkyl, or salt cation;

a compound having the formula:

$$R_{1} = O - C - COOR_{9}$$

$$R_{1} = O - C - COOR_{9}$$

where $R_{3\cdot6}$ can be the hydrogen, halogen, substituted or unsubstituted $C_{1\cdot3}$ alkyl group, or a $C_{1\cdot3}$ ether or ester, and these moieties may be the same or different, or alkyl moieties of an aromatic or aliphatic ring incorporating two of the $R_{3\cdot6}$,

R₁ can be connected to any position on the phenyl ring, and

sites $R_{7.8}$ can be hydrogen, halogen, methyl, ethyl, and these moieties may be the same or different, or alkyl groups in a ring connecting the two, and

R₉ can be a hydrogen, halogen, C₁₋₃ lower alkyl, or salt cation;

a compound having the formula:

where R1 can be connected to any position on the phenyl ring, and

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sites $R_{7.8}$ can be hydrogen, halogen, methyl, ethyl, and these moieties may be the same or different, or alkyl groups in a ring connecting the two, and

 R_2 is defined as a substituted or unsubstituted aromatic compound, a substituted or unsubstituted alkyl ring compound, or a substituted or unsubstituted phthalimide compound,

X is a carboxyl,

Y is a nitrogen,

and R_2 completes the phthalimide compound by being bonded to both X and Y; and where X, Y, and Z, may either be CH_2 , NH, O, or N, with the caveat that each are different from the other;

a compound having the formula:

$$R_3$$
 R_4
 R_6
 R_7
 R_8
 R_7
 R_8
 R_7
 R_8

where R2, R3, R4, R5, and R6 may be hydrogen, halogen, or alkyl groups and may be the same or different.

 $R_{\rm 7}$ and $R_{\rm 8}$ may be hydrogen or methyl groups and may be the same or different, and where the $R_{\rm 9}$ moiety is hydrogen or a salt cation;

a compound having the formula:

where R_2 is a substituted or unsubstituted aromatic compound, or a substituted or unsubstituted alkyl ring compound, or a substituted or unsubstituted phthalimide compound that incorporates X and Y,

X is a carbonyl,

Y is a nitrogen, and

 R_2 completes the phthalimide compound by being bonded to both X and Y, and where X, Y, and Z are CH_2 , NH, S, SO_2 , CO, O or N with the caveat that X, Y, and Z are each different from one another, and

where R1 can be connected to any position on the phenyl ring, and

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R₃ and R₄ are hydrogen, halogen, methyl, ethyl, propyl, isopropyl, neopentyl, butyl, or substituted or unsubstituted aryl groups and these moieties may be the same or different, or alkyl moieties as part of an aliphatic ring connecting R3 and R4, and

R₅ is a hydrogen, halogen, C₁₋₃ lower alkyl, or a salt cation; a compound having the formula:

where A is a chemical bridge which includes two to four chemical moieties bonded together,

the chemical moieties in A are selected from the group consisting of CO, O, S, SO₂, NH, NR9 where R9 is a C1-6 alkyl group, CH2, CH, and C, with the proviso that, except in the case where A contains two identical CH and C moieties positioned adjacent one another to form an alkene or alkyne, the chemical moieties in A are each different from one another, and at least one of R₁₋₅ is substituted with a compound having the chemical formula:

$$O-(CH_2)_n$$
 $-C-COOR_{12}$

where n is zero to five,

where R_{10} and R_{11} are selected from the group consisting of hydrogen, halogen, $C_{1\text{-}12}$ alkyl groups, carboxylic acids and esters, aromatic or heteroatomic groups, and these moieties may be the same or different, or alkyl moieties of part of an aliphatic ring connecting R₁₀ and R11, and where R12 is a hydrogen, halogen, salt cation, metal, or C1-6 alkyl group, and wherein a remainder of the R1-5 moieties and the R6-8 moieties are selected from the group consisting of hydrogen, halogen, C1-6 alkyl groups, C1-6 ether or esters, aromatics and heteroaromatics, and alkyl moieties of an aliphatic ring connecting two sites on a phenyl group; a compound having the formula:

$$R_1$$
— A — R_2

compounds, or a substituted or unsubstituted alkyl or heteroalkyl ring compound, or a substituted or unsubstituted phthalimide compound, and

where R1 and R2 may be the same or different,

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where A is a chemical bridge which includes three chemical moieties bonded together between R₁ and R₂.

wherein the chemical moieties in A are selected from the group consisting of CO, O, S, SO_2 , NH, NR_3 where R_3 is C_{1-6} alkyl group, NR_4 where R_4 includes two carbonyls as part of a phthalimide compound formed with R_1 or R_2 , CH_2 , CH, and C, and

where at least one of R_1 and R_2 is substituted with a compounds having the chemical formula:

$$O-(CH_2)_n$$
 C
 $COOR_7$
 R_6

where n is zero to five, where R_5 and R_6 are selected from the group consisting of hydrogen, halogen, substituted or unsubstituted C_{1-12} alkyl groups, carboxylic acid and ester groups, substituted or unsubstituted aromatic or heteroaromatic groups, and these moieties may be the same or different, or alkyl moieties of part of an aliphatic ring connecting R_5 and R_6 , and

where R_7 is a hydrogen, halogen, salt cation, metal, or substituted or unsubstituted $C_{1.6}$ alkyl group;

a compound having the formula:

$$R_1$$
— A — R_2

where R_1 and R_2 each are a substituted or unsubstituted aromatic or heteroaromatic compound, or substituted or unsubstituted alkyl or heteroalkyl ring compound, or a substituted or unsubstituted phthalimide compound, and

where R1 and R2 may be the same or different,

where A is a chemical bridge which includes two to four chemical moieties bonded together between R₁ and R₂,

wherein said chemical moieties in A are selected from the group consisting of CO, O, S, SO_2 , NH, NR_3 where R_3 is a C_{1-6} alkyl group, NR_4 where R_4 includes two carbonyls as part of a phthalimide compound formed with R_1 or R_2 , CH_2 , CH, and C, with the caveat that, except in the case where A contains two identical CH and C moieties positioned adjacent one another to form an alkene or alkyne, the chemical moieties in A are each different from one another, and

wherein at least one of R_1 or R_2 is substituted with a compound having the chemical formula:

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where n is zero to five,

where R_5 and R_6 are selected from the group consisting of hydrogen, halogen, substituted or unsubstituted C_{1-12} alkyl groups, carboxylic acid and ester, substituted or unsubstituted aromatic or heteroaromatic groups, and these moieties may be the same or different, or alkyl moieties of part of an aliphatic ring connecting R_5 and R_6 , and

where R_7 is a hydrogen, halogen, salt cation, metal, or substituted or unsubstituted $C_{1\cdot 6}$ alkyl group; and

a compound having the formula:

$$\begin{array}{c|c} & H & O & H \\ \hline & H & O & H \\ \hline & R_1 - N - C - C & - C \\ \hline & H & R_3 \end{array}$$

where R_1 is selected from the group consisting of optionally substituted phenyl, adamantyl, napthyl, and indanyl, $R_{2\cdot 3}$ are alkyl moieties of a $C_{3\cdot 6}$ alkyl ring connecting R_2 and R_3 , and R_4 is a hydrogen, a monovalent salt cation, or a $C_{1\cdot 3}$ lower alkyl.

- The method of Claim 8, wherein the allosteric effector compound is administered at a dose of 100-300 mg/kg.
- The method of Claim 7, wherein the allosteric effector compound is 2-[4-(((3,5-dimethylanilino)carbonyl)methyl)phenoxy]-2-methylpropionic acid, or a physiologically acceptable salt thereof.
- The method of Claim 10, wherein the allosteric effector compound is administered at a dose of 100-300 mg/kg.
- The method of Claim 10, wherein the allosteric effector compound is administered at a dose of 200 mg/kg.
 - 13. A method of measuring tumor oxygenation, comprising

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- a) administering an effective amount of an allosteric effector compound capable of decreasing hemoglobin binding affinity for oxygen; and
- b) performing a blood oxygen level-dependent magnetic resonance imaging scan, whereby oxygenation of the tumor is measured.
- 14. The method of Claim 13, wherein the allosteric effector compound is selected from the group consisting of:

a compound having the formula:

where $R_{1.5}$ may be hydrogen, halogen, or a substituted or unsubstituted $C_{1.3}$ alkyl group and may be the same or different,

R₆₋₇ may each be hydrogen or methyl and may be the same or different, and

 R_8 may be hydrogen, a substituted or unsubstituted $C_{1\cdot3}$ alkyl group, or a salt cation, and X and Z are CH_2 , NH, or O;

a compound having the formula:

$$R_4$$
 R_5
 R_6
 R_6
 R_7
 R_7
 R_7
 R_8

where X and Z may each be CH₂, CO, NH or O, and Y may be CO or NH, which the caveat that X, Y, and Z must all be different from each other, and

 R_{2-6} can be the hydrogen, halogen, substituted or unsubstituted C_{1-3} alkyl groups, and may be the same or different,

 $R_{7.8}$ can be hydrogens, methyls, ethyls, or alkyl groups in a ring connecting the two, and R_0 can be a hydrogen, lower alkyl, or salt cation;

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where $R_{3\cdot6}$ can be the hydrogen, halogen, substituted or unsubstituted $C_{1\cdot3}$ alkyl group, or a $C_{1\cdot3}$ ether or ester, and these moieties may be the same or different, or alkyl moieties of an aromatic or aliphatic ring incorporating two of the $R_{3\cdot6}$,

R₁ can be connected to any position on the phenyl ring, and

sites $R_{7.8}$ can be hydrogen, halogen, methyl, ethyl, and these moieties may be the same or different, or alkyl groups in a ring connecting the two, and

R₉ can be a hydrogen, halogen, C₁₋₃ lower alkyl, or salt cation; a compound having the formula:

$$R_2 = X$$

$$R_1 = 0 - C - COOR_{\epsilon}$$

$$R_2 = 0 - C - COOR_{\epsilon}$$

where R1 can be connected to any position on the phenyl ring, and

sites $R_{7.8}$ can be hydrogen, halogen, methyl, ethyl, and these moieties may be the same or different, or alkyl groups in a ring connecting the two, and

 R_2 is defined as a substituted or unsubstituted aromatic compound, a substituted or unsubstituted alkyl ring compound, or a substituted or unsubstituted phthalimide compound,

X is a carboxyl,

Y is a nitrogen,

and R_2 completes the phthalimide compound by being bonded to both X and Y; and where X, Y, and Z, may either be CH_2 , NH, O, or N, with the caveat that each are different from the other:

$$R_3$$
 R_4
 R_6
 R_7
 R_8
 R_7
 R_8
 R_8

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where R_2 , R_3 , R_4 , R_5 , and R_6 may be hydrogen, halogen, or alkyl groups and may be the same or different.

 R_7 and R_8 may be hydrogen or methyl groups and may be the same or different, and where the R_9 moiety is hydrogen or a salt cation;

a compound having the formula:

where R_2 is a substituted or unsubstituted aromatic compound, or a substituted or unsubstituted alkyl ring compound, or a substituted or unsubstituted phthalimide compound that incorporates X and Y,

X is a carbonyl,

Y is a nitrogen, and

R₂ completes the phthalimide compound by being bonded to both X and Y, and where X, Y, and Z are CH₂, NH, S, SO₂, CO, O or N with the caveat that X, Y, and Z are each different from one another, and

where R1 can be connected to any position on the phenyl ring, and

R₃ and R₄ are hydrogen, halogen, methyl, ethyl, propyl, isopropyl, neopentyl, butyl, or substituted or unsubstituted aryl groups and these moieties may be the same or different, or alkyl moieties as part of an aliphatic ring connecting R₃ and R₄, and

R₅ is a hydrogen, halogen, C₁₋₃ lower alkyl, or a salt cation; a compound having the formula:

where A is a chemical bridge which includes two to four chemical moieties bonded together,

the chemical moieties in A are selected from the group consisting of CO, O, S, SO₂, NH, NR₉ where R₉ is a C₁₋₆ alkyl group, CH₂, CH, and C, with the proviso that, except in the case where A contains two identical CH and C moieties positioned adjacent one another to form an alkene or alkyne, the chemical moieties in A are each different from one another, and

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at least one of R1-5 is substituted with a compound having the chemical formula:

$$O-(CH_2)_n$$
 $C-COOR_{12}$ R_{11}

where n is zero to five.

where R_{10} and R_{11} are selected from the group consisting of hydrogen, halogen, C_{1-12} alkyl groups, carboxylic acids and esters, aromatic or heteroatomic groups, and these moieties may be the same or different, or alkyl moieties of part of an aliphatic ring connecting R_{10} and R_{11} , and where R_{12} is a hydrogen, halogen, salt cation, metal, or C_{1-6} alkyl group, and wherein a remainder of the R_{1-5} moieties and the R_{6-8} moieties are selected from the group consisting of hydrogen, halogen, C_{1-6} alkyl groups, C_{1-6} ether or esters, aromatics and heteroaromatics, and alkyl moieties of an aliphatic ring connecting two sites on a phenyl group; a compound having the formula:

where R_1 and R_2 each are a substituted or unsubstituted aromatic or heteroaromatic compounds, or a substituted or unsubstituted alkyl or heteroalkyl ring compound, or a substituted or unsubstituted phthalimide compound, and

where R1 and R2 may be the same or different,

where A is a chemical bridge which includes three chemical moieties bonded together between R_1 and R_2 ,

wherein the chemical moieties in A are selected from the group consisting of CO, O, S, SO₂, NH, NR₃ where R₃ is C₁₋₆ alkyl group, NR₄ where R₄ includes two carbonyls as part of a phthalimide compound formed with R₁ or R₂, CH₂, CH, and C, and

where at least one of R₁ and R₂ is substituted with a compounds having the chemical formula:

where n is zero to five, where R_5 and R_6 are selected from the group consisting of hydrogen, halogen, substituted or unsubstituted C_{1-12} alkyl groups, carboxylic acid and ester groups, substituted or unsubstituted aromatic or heteroaromatic groups, and these moieties may be the same or different, or alkyl moieties of part of an aliphatic ring connecting R_5 and R_6 , and

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where R_7 is a hydrogen, halogen, salt cation, metal, or substituted or unsubstituted $C_{1.6}$ alkyl group;

a compound having the formula:

$$R_1 - A - R_2$$

where R_1 and R_2 each are a substituted or unsubstituted aromatic or heteroaromatic compound, or substituted or unsubstituted alkyl or heteroalkyl ring compound, or a substituted or unsubstituted phthalimide compound, and

where R1 and R2 may be the same or different,

where A is a chemical bridge which includes two to four chemical moieties bonded together between R_1 and R_2 ,

wherein said chemical moieties in A are selected from the group consisting of CO, O, S, SO_2 , NH, NR_3 where R_3 is a $C_{1\cdot6}$ alkyl group, NR_4 where R_4 includes two carbonyls as part of a phthalimide compound formed with R_1 or R_2 , CH, and C, with the caveat that, except in the case where A contains two identical CH and C moieties positioned adjacent one another to form an alkene or alkyne, the chemical moieties in A are each different from one another, and

wherein at least one of R_1 or R_2 is substituted with a compound having the chemical formula:

$$O-(CH_2)_n$$
 $-C-COOR_7$

where n is zero to five,

where R_5 and R_6 are selected from the group consisting of hydrogen, halogen, substituted or unsubstituted C_{1-12} alkyl groups, carboxylic acid and ester, substituted or unsubstituted aromatic or heteroaromatic groups, and these moieties may be the same or different, or alkyl moieties of part of an aliphatic ring connecting R_5 and R_6 , and

where R_7 is a hydrogen, halogen, salt cation, metal, or substituted or unsubstituted $C_{1.6}$ alkyl group; and

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where R_1 is selected from the group consisting of optionally substituted phenyl, adamantyl, napthyl, and indanyl, $R_{2\cdot 3}$ are alkyl moieties of a $C_{3\cdot 6}$ alkyl ring connecting R_2 and R_3 , and R_4 is a hydrogen, a monovalent salt cation, or a $C_{1\cdot 3}$ lower alkyl.

- The method of Claim 14, wherein the allosteric effector compound is administered at a dose of 100-300 mg/kg.
- The method of Claim 13, wherein the oxygenation of the tumor is measured quantitatively.
- The method of Claim 13, wherein the allosteric effector compound is 2-[4-(((3,5-dimethylanilino)carbonyl)methyl)phenoxy]-2-methylpropionic acid, or a physiologically acceptable salt thereof.
- The method of Claim 17, wherein the allosteric effector compound is administered at a dose of 100-300 mg/kg.
- The method of Claim 17, wherein the allosteric effector compound is administered at a dose of 200 mg/kg.
- A method for determining an optimal time for the initiation of radiation therapy, comprising
- a) administering an effective amount of an allosteric effector compound capable of decreasing hemoglobin binding affinity for oxygen;
 - b) measuring the blood oxygen level dependent signal ratio; and
- c) determining the time required for a maximum increase in the blood oxygen level dependent signal ratio, whereby the optimal time for initiation of radiation therapy is determined.
- 21. The method of Claim 20, wherein the allosteric effector compound is selected from the group consisting of:
 - a compound having the formula:

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where $R_{1:5}$ may be hydrogen, halogen, or a substituted or unsubstituted $C_{1:3}$ alkyl group and may be the same or different,

R₆₋₇ may each be hydrogen or methyl and may be the same or different, and

 R_8 may be hydrogen, a substituted or unsubstituted C_{1-3} alkyl group, or a salt cation, and X and Z are CH_2 , NH, or O;

a compound having the formula:

where X and Z may each be CH₂, CO, NH or O, and Y may be CO or NH, which the caveat that X, Y, and Z must all be different from each other, and

 $R_{2.6}$ can be the hydrogen, halogen, substituted or unsubstituted $C_{1.3}$ alkyl groups, and may be the same or different,

 R_{7-8} can be hydrogens, methyls, ethyls, or alkyl groups in a ring connecting the two, and R_9 can be a hydrogen, lower alkyl, or salt cation;

a compound having the formula:

$$R_{1}$$
 R_{1} R_{1} R_{1} R_{1} R_{1} R_{2} R_{3} R_{4} R_{5} R_{6} R_{6}

where R_{3-6} can be the hydrogen, halogen, substituted or unsubstituted C_{1-3} alkyl group, or a C_{1-3} ether or ester, and these moieties may be the same or different, or alkyl moieties of an aromatic or aliphatic ring incorporating two of the R_{3-6} ,

R₁ can be connected to any position on the phenyl ring, and

sites $R_{7.8}$ can be hydrogen, halogen, methyl, ethyl, and these moieties may be the same or different, or alkyl groups in a ring connecting the two, and

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 R_9 can be a hydrogen, halogen, C_{1-3} lower alkyl, or salt cation; a compound having the formula:

$$R_2 - X$$

$$Z$$

$$R_1 = O - C - COOR_9$$

$$R_8$$

where R1 can be connected to any position on the phenyl ring, and

sites $R_{7.8}$ can be hydrogen, halogen, methyl, ethyl, and these moieties may be the same or different, or alkyl groups in a ring connecting the two, and

 R_2 is defined as a substituted or unsubstituted aromatic compound, a substituted or unsubstituted alkyl ring compound, or a substituted or unsubstituted phthalimide compound,

X is a carboxyl,

Y is a nitrogen,

and R_2 completes the phthalimide compound by being bonded to both X and Y; and where X, Y, and Z, may either be CH_2 , NH, O, or N, with the caveat that each are different from the other:

a compound having the formula:

$$\begin{array}{c|c} R_3 & R_2 & O \\ R_4 & R_5 & R_6 & COOR_{\epsilon} \end{array}$$

where R2, R3, R4, R5, and R6 may be hydrogen, halogen, or alkyl groups and may be the same or different.

 R_7 and R_8 may be hydrogen or methyl groups and may be the same or different, and where the R_9 moiety is hydrogen or a salt cation;

a compound having the formula:

where R_2 is a substituted or unsubstituted aromatic compound, or a substituted or unsubstituted alkyl ring compound, or a substituted or unsubstituted phthalimide compound that incorporates X and Y,

X is a carbonyl,

Y is a nitrogen, and

R₂ completes the phthalimide compound by being bonded to both X and Y, and where X, Y, and Z are CH₂, NH, S, SO₂, CO, O or N with the caveat that X, Y, and Z are each different from one another, and

where R₁ can be connected to any position on the phenyl ring, and

R₃ and R₄ are hydrogen, halogen, methyl, ethyl, propyl, isopropyl, neopentyl, butyl, or substituted or unsubstituted aryl groups and these moieties may be the same or different, or alkyl moieties as part of an aliphatic ring connecting R₃ and R₄, and

 R_5 is a hydrogen, halogen, C_{1-3} lower alkyl, or a salt cation; a compound having the formula:

where A is a chemical bridge which includes two to four chemical moieties bonded together,

the chemical moieties in A are selected from the group consisting of CO, O, S, SO_2 , NH, NR_9 where R_9 is a C_{1-6} alkyl group, CH_2 , CH, and C, with the proviso that, except in the case where A contains two identical CH and C moieties positioned adjacent one another to form an alkene or alkyne, the chemical moieties in A are each different from one another, and

at least one of R₁₋₅ is substituted with a compound having the chemical formula:

where n is zero to five,

where R_{10} and R_{11} are selected from the group consisting of hydrogen, halogen, $C_{1\cdot 12}$ alkyl groups, carboxylic acids and esters, aromatic or heteroatomic groups, and these moieties may be the same or different, or alkyl moieties of part of an aliphatic ring connecting R_{10} and R_{11} , and where R_{12} is a hydrogen, halogen, salt cation, metal, or $C_{1\cdot 6}$ alkyl group, and wherein a remainder of the $R_{1\cdot 5}$ moieties and the $R_{6\cdot 8}$ moieties are selected from the group consisting of hydrogen, halogen, $C_{1\cdot 6}$ alkyl groups, $C_{1\cdot 6}$ ether or esters, aromatics and heteroaromatics, and alkyl moieties of an aliphatic ring connecting two sites on a phenyl group;

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a compound having the formula:

where R_1 and R_2 each are a substituted or unsubstituted aromatic or heteroaromatic compounds, or a substituted or unsubstituted alkyl or heteroalkyl ring compound, or a substituted or unsubstituted phthalimide compound, and

where R₁ and R₂ may be the same or different,

where A is a chemical bridge which includes three chemical moieties bonded together between R₁ and R₂.

wherein the chemical moieties in A are selected from the group consisting of CO, O, S, SO₂, NH, NR₃ where R₃ is C₁₋₆ alkyl group, NR₄ where R₄ includes two carbonyls as part of a phthalimide compound formed with R₁ or R₂, CH₂, CH, and C, and

where at least one of R_1 and R_2 is substituted with a compounds having the chemical formula:

where n is zero to five, where R_5 and R_6 are selected from the group consisting of hydrogen, halogen, substituted or unsubstituted C_{1-12} alkyl groups, carboxylic acid and ester groups, substituted or unsubstituted aromatic or heteroaromatic groups, and these moieties may be the same or different, or alkyl moieties of part of an aliphatic ring connecting R_5 and R_6 , and

where R_7 is a hydrogen, halogen, salt cation, metal, or substituted or unsubstituted C_{1-6} alkyl group;

a compound having the formula:

where R_1 and R_2 each are a substituted or unsubstituted aromatic or heteroaromatic compound, or substituted or unsubstituted alkyl or heteroalkyl ring compound, or a substituted or unsubstituted phthalimide compound, and

where R1 and R2 may be the same or different,

where A is a chemical bridge which includes two to four chemical moieties bonded together between R_1 and R_2 ,

wherein said chemical moieties in A are selected from the group consisting of CO, O, S, SO_2 , NH, NR₃ where R_3 is a C_{1-6} alkyl group, NR₄ where R_4 includes two carbonyls as part of a phthalimide compound formed with R_1 or R_2 , CH₂, CH, and C, with the caveat that, except in the

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case where A contains two identical CH and C moieties positioned adjacent one another to form an alkene or alkyne, the chemical moieties in A are each different from one another, and

wherein at least one of R_1 or R_2 is substituted with a compound having the chemical formula:

$$O-(CH_2)_n$$
 $-C$
 $-COOR_7$
 R_6

where n is zero to five.

where R_5 and R_6 are selected from the group consisting of hydrogen, halogen, substituted or unsubstituted C_{1-12} alkyl groups, carboxylic acid and ester, substituted or unsubstituted aromatic or heteroaromatic groups, and these moieties may be the same or different, or alkyl moieties of part of an aliphatic ring connecting R_5 and R_6 , and

where R_7 is a hydrogen, halogen, salt cation, metal, or substituted or unsubstituted $C_{1.6}$ alkyl group; and

a compound having the formula:

where R_1 is selected from the group consisting of optionally substituted phenyl, adamantyl, napthyl, and indanyl, $R_{2\cdot 3}$ are alkyl moieties of a $C_{3\cdot 6}$ alkyl ring connecting R_2 and R_3 , and R_4 is a hydrogen, a monovalent salt cation, or a $C_{1\cdot 3}$ lower alkyl.

- 22. The method of Claim 20, wherein the allosteric effector compound is 2-[4-(((3,5-dimethylanilino)carbonyl)methyl)phenoxy]-2-methylpropionic acid, or a physiologically acceptable salt thereof.
 - A method of diagnosing an abnormal pathology, comprising
- a) introducing an allosteric effector compound into a patient suspected of having the abnormal pathology;
- b) performing a blood oxygen level-dependent magnetic resonance imaging scan of the patient, and
- c) detecting an increase in blood oxygen level-dependent magnetic resonance imaging signal; whereby an abnormal pathology may be diagnosed.

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24. The method of Claim 23, wherein the allosteric effector compound is selected from the group consisting of:

a compound having the formula:

$$R_4$$
 R_5
 R_6
 R_6
 R_7
 R_6
 R_6

where R_{1-5} may be hydrogen, halogen, or a substituted or unsubstituted C_{1-3} alkyl group and may be the same or different,

 R_{6-7} may each be hydrogen or methyl and may be the same or different, and

 R_8 may be hydrogen, a substituted or unsubstituted C_{1-3} alkyl group, or a salt cation, and X and Z are CH_2 , NH, or O;

a compound having the formula:

$$R_3$$
 R_2
 R_4
 R_5
 R_6
 R_6
 R_6

where X and Z may each be CH₂, CO, NH or O, and Y may be CO or NH, which the caveat that X, Y, and Z must all be different from each other, and

 $R_{2.6}$ can be the hydrogen, halogen, substituted or unsubstituted $C_{1.3}$ alkyl groups, and may be the same or different,

 R_{7-8} can be hydrogens, methyls, ethyls, or alkyl groups in a ring connecting the two, and R_9 can be a hydrogen, lower alkyl, or salt cation;

a compound having the formula:

$$R_{4}$$

$$R_{5}$$

$$R_{6}$$

$$R_{6}$$

$$R_{1} = 0 - C - COOR_{4}$$

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where R_{3-6} can be the hydrogen, halogen, substituted or unsubstituted C_{1-3} alkyl group, or a C_{1-3} ether or ester, and these moieties may be the same or different, or alkyl moieties of an aromatic or aliphatic ring incorporating two of the R_{3-6} ,

R₁ can be connected to any position on the phenyl ring, and

sites $R_{7.8}$ can be hydrogen, halogen, methyl, ethyl, and these moieties may be the same or different, or alkyl groups in a ring connecting the two, and

 R_9 can be a hydrogen, halogen, $C_{1\text{-}3}$ lower alkyl, or salt cation; a compound having the formula:

where R1 can be connected to any position on the phenyl ring, and

sites $R_{7.8}$ can be hydrogen, halogen, methyl, ethyl, and these moieties may be the same or different, or alkyl groups in a ring connecting the two, and

 $\rm R_2$ is defined as a substituted or unsubstituted aromatic compound, a substituted or unsubstituted alkyl ring compound, or a substituted or unsubstituted phthalimide compound,

X is a carboxyl,

Y is a nitrogen,

and R_2 completes the phthalimide compound by being bonded to both X and Y; and where X, Y, and Z, may either be CH_2 , NH, O, or N, with the caveat that each are different from the other:

a compound having the formula:

$$R_3$$
 R_4
 R_6
 R_7
 R_8
 R_7
 R_8
 R_8

where R_2 , R_3 , R_4 , R_5 , and R_6 may be hydrogen, halogen, or alkyl groups and may be the same or different.

 R_7 and R_8 may be hydrogen or methyl groups and may be the same or different, and where the R_9 moiety is hydrogen or a salt cation;

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where R_2 is a substituted or unsubstituted aromatic compound, or a substituted or unsubstituted alkyl ring compound, or a substituted or unsubstituted phthalimide compound that incorporates X and Y,

X is a carbonyl,

Y is a nitrogen, and

R₂ completes the phthalimide compound by being bonded to both X and Y, and where X, Y, and Z are CH₂, NH, S, SO₂, CO, O or N with the caveat that X, Y, and Z are each different from one another, and

where R1 can be connected to any position on the phenyl ring, and

R₃ and R₄ are hydrogen, halogen, methyl, ethyl, propyl, isopropyl, neopentyl, butyl, or substituted or unsubstituted aryl groups and these moieties may be the same or different, or alkyl moieties as part of an aliphatic ring connecting R₃ and R₄, and

R₅ is a hydrogen, halogen, C₁₋₃ lower alkyl, or a salt cation; a compound having the formula:

where A is a chemical bridge which includes two to four chemical moieties bonded together,

the chemical moieties in A are selected from the group consisting of CO, O, S, SO_2 , NH, NR₉ where R₉ is a C₁₋₆ alkyl group, CH₂, CH, and C, with the proviso that, except in the case where A contains two identical CH and C moieties positioned adjacent one another to form an alkene or alkyne, the chemical moieties in A are each different from one another, and

at least one of R₁₋₅ is substituted with a compound having the chemical formula:

$$O-(CH_2)_n$$
 $C-COOR_{12}$

25 where n is zero to five,

alkyl group;

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where R_{10} and R_{11} are selected from the group consisting of hydrogen, halogen, C_{1-12} alkyl groups, carboxylic acids and esters, aromatic or heteroatomic groups, and these moieties may be the same or different, or alkyl moieties of part of an aliphatic ring connecting R_{10} and R_{11} , and where R_{12} is a hydrogen, halogen, salt cation, metal, or C_{1-6} alkyl group, and wherein a remainder of the R_{1-5} moieties and the R_{6-8} moieties are selected from the group consisting of hydrogen, halogen, C_{1-6} alkyl groups, C_{1-6} ether or esters, aromatics and heteroaromatics, and alkyl moieties of an aliphatic ring connecting two sites on a phenyl group;

a compound having the formula:

where R_1 and R_2 each are a substituted or unsubstituted aromatic or heteroaromatic compounds, or a substituted or unsubstituted alkyl or heteroalkyl ring compound, or a substituted or unsubstituted phthalimide compound, and

where R1 and R2 may be the same or different,

where A is a chemical bridge which includes three chemical moieties bonded together between R_1 and R_2 ,

wherein the chemical moieties in A are selected from the group consisting of CO, O, S, SO₂, NH, NR₃ where R_3 is C_{1-6} alkyl group, NR₄ where R_4 includes two carbonyls as part of a phthalimide compound formed with R_1 or R_2 , CH₂, CH, and C, and

where at least one of \mathbf{R}_1 and \mathbf{R}_2 is substituted with a compounds having the chemical formula:

where n is zero to five, where R_5 and R_6 are selected from the group consisting of hydrogen, halogen, substituted or unsubstituted $C_{1\text{-}12}$ alkyl groups, carboxylic acid and ester groups, substituted or unsubstituted aromatic or heteroaromatic groups, and these moieties may be the same or different, or alkyl moieties of part of an aliphatic ring connecting R_5 and R_6 , and where R_7 is a hydrogen, halogen, salt cation, metal, or substituted or unsubstituted $C_{1\text{-}6}$

a compound having the formula:

$$R_1 - A - R_2$$

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where R_1 and R_2 each are a substituted or unsubstituted aromatic or heteroaromatic compound, or substituted or unsubstituted alkyl or heteroalkyl ring compound, or a substituted or unsubstituted phthalimide compound, and

where R1 and R2 may be the same or different.

where A is a chemical bridge which includes two to four chemical moieties bonded together between R_1 and R_2 ,

wherein said chemical moieties in A are selected from the group consisting of CO, O, S, SO_2 , NH, NR_3 where R_3 is a C_{1-6} alkyl group, NR_4 where R_4 includes two carbonyls as part of a phthalimide compound formed with R_1 or R_2 , CH_2 , CH, and C, with the caveat that, except in the case where A contains two identical CH and C moieties positioned adjacent one another to form an alkene or alkyne, the chemical moieties in A are each different from one another, and

wherein at least one of R_1 or R_2 is substituted with a compound having the chemical formula:

$$O-(CH_2)_{ri}-C-COOR_7$$

where n is zero to five,

where R_5 and R_6 are selected from the group consisting of hydrogen, halogen, substituted or unsubstituted C_{1-12} alkyl groups, carboxylic acid and ester, substituted or unsubstituted aromatic or heteroaromatic groups, and these moieties may be the same or different, or alkyl moieties of part of an aliphatic ring connecting R_5 and R_6 , and

where R_7 is a hydrogen, halogen, salt cation, metal, or substituted or unsubstituted $C_{1.6}$ alkyl group; and

a compound having the formula:

where R_1 is selected from the group consisting of optionally substituted phenyl, adamantyl, napthyl, and indanyl, $R_{2\cdot3}$ are alkyl moieties of a $C_{3\cdot6}$ alkyl ring connecting R_2 and R_3 , and R_4 is a hydrogen, a monovalent salt cation, or a $C_{1\cdot3}$ lower alkyl.

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- The method of Claim 23, wherein the allosteric effector compound is 2-[4-(((3,5-dimethylanilino)carbonyl)methyl)phenoxy]-2-methylpropionic acid, or a physiologically acceptable salt thereof.
 - 26. A method of imaging glioblastoma multiforme, comprising
- a) administering an effective amount of an allosteric effector compound capable of decreasing hemoglobin binding affinity for oxygen; and
- b) performing a blood oxygen level-dependent magnetic resonance imaging scan, whereby glioblastoma multifome may be imaged.
- 27. The method of Claim 26, wherein the allosteric effector compound is selected from the group consisting of:

a compound having the formula:

where $R_{1.5}$ may be hydrogen, halogen, or a substituted or unsubstituted $C_{1.3}$ alkyl group and may be the same or different,

R₆₋₇ may each be hydrogen or methyl and may be the same or different, and

 R_8 may be hydrogen, a substituted or unsubstituted $C_{1\text{-}3}$ alkyl group, or a salt cation, and X and Z are CH₂, NH, or O;

a compound having the formula:

where X and Z may each be CH₂, CO, NH or O, and Y may be CO or NH, which the caveat that X, Y, and Z must all be different from each other, and

 $R_{2.6}$ can be the hydrogen, halogen, substituted or unsubstituted $C_{1.3}$ alkyl groups, and may be the same or different,

 $R_{7.8}$ can be hydrogens, methyls, ethyls, or alkyl groups in a ring connecting the two, and R_9 can be a hydrogen, lower alkyl, or salt cation;

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a compound having the formula:

$$R_4$$

$$R_6$$

$$R_1 = 0$$

$$R_8$$

$$R_8$$

where R_{3-6} can be the hydrogen, halogen, substituted or unsubstituted C_{1-3} alkyl group, or a C_{1-3} ether or ester, and these moieties may be the same or different, or alkyl moieties of an aromatic or aliphatic ring incorporating two of the R_{3-6} ,

R1 can be connected to any position on the phenyl ring, and

sites $R_{7.8}$ can be hydrogen, halogen, methyl, ethyl, and these moieties may be the same or different, or alkyl groups in a ring connecting the two, and

 $R_{\rm 9}$ can be a hydrogen, halogen, $C_{\rm 1-3}$ lower alkyl, or salt cation; a compound having the formula:

$$R_2 = X$$

$$R_1 = O - C - COOR_9$$

$$R_8$$

where R1 can be connected to any position on the phenyl ring, and

sites $R_{7.8}$ can be hydrogen, halogen, methyl, ethyl, and these moieties may be the same or different, or alkyl groups in a ring connecting the two, and

 R_2 is defined as a substituted or unsubstituted aromatic compound, a substituted or unsubstituted alkyl ring compound, or a substituted or unsubstituted phthalimide compound,

X is a carboxyl,

Y is a nitrogen,

and R_2 completes the phthalimide compound by being bonded to both X and Y; and where X, Y, and Z, may either be CH_2 , NH, O, or N, with the caveat that each are different from the other;

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$$R_3$$
 R_4
 R_5
 R_6
 R_7
 R_8
 R_7
 R_8
 R_9
 R_9

where R_2 , R_3 , R_4 , R_5 , and R_6 may be hydrogen, halogen, or alkyl groups and may be the same or different.

 R_7 and R_8 may be hydrogen or methyl groups and may be the same or different, and where the R_9 moiety is hydrogen or a salt cation;

a compound having the formula:

where R_2 is a substituted or unsubstituted aromatic compound, or a substituted or unsubstituted alkyl ring compound, or a substituted or unsubstituted phthalimide compound that incorporates X and Y,

X is a carbonyl,

Y is a nitrogen, and

 R_2 completes the phthalimide compound by being bonded to both X and Y, and where X, Y, and Z are CH_2 , NH, S, SO_2 , CO, O or N with the caveat that X, Y, and Z are each different from one another, and

where R1 can be connected to any position on the phenyl ring, and

R₃ and R₄ are hydrogen, halogen, methyl, ethyl, propyl, isopropyl, neopentyl, butyl, or substituted or unsubstituted aryl groups and these moieties may be the same or different, or alkyl moieties as part of an aliphatic ring connecting R₃ and R₄, and

 R_5 is a hydrogen, halogen, C_{1-3} lower alkyl, or a salt cation; a compound having the formula:

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where A is a chemical bridge which includes two to four chemical moieties bonded together.

the chemical moieties in A are selected from the group consisting of CO, O, S, SO2, NH, NR₀ where R₀ is a C₁₋₆ alkyl group, CH₂, CH, and C, with the proviso that, except in the case where A contains two identical CH and C moieties positioned adjacent one another to form an alkene or alkyne, the chemical mojeties in A are each different from one another, and

at least one of R₁₋₅ is substituted with a compound having the chemical formula:

$$O-(CH_2)_n$$
 $C-COOR_{12}$

where n is zero to five,

where R₁₀ and R₁₁ are selected from the group consisting of hydrogen, halogen, C₁₋₁₂ alkyl groups, carboxylic acids and esters, aromatic or heteroatomic groups, and these moieties may be the same or different, or alkyl moieties of part of an aliphatic ring connecting R10 and R11, and where R12 is a hydrogen, halogen, salt cation, metal, or C1-6 alkyl group, and wherein a remainder of the R₁₋₅ moieties and the R₆₋₈ moieties are selected from the group consisting of hydrogen, halogen, C1-6 alkyl groups, C1-6 ether or esters, aromatics and heteroaromatics, and alkyl moieties of an aliphatic ring connecting two sites on a phenyl group; a compound having the formula:

 $R_1\!\!-\!\!A\!\!-\!\!R_2$ where R_1 and R_2 each are a substituted or unsubstituted aromatic or heteroaromatic compounds, or a substituted or unsubstituted alkyl or heteroalkyl ring compound, or a substituted or unsubstituted phthalimide compound, and

where R1 and R2 may be the same or different,

where A is a chemical bridge which includes three chemical moieties bonded together between R1 and R2,

wherein the chemical moieties in A are selected from the group consisting of CO, O, S, SO₂, NH, NR₃ where R₃ is C₁₋₆ alkyl group, NR₄ where R₄ includes two carbonyls as part of a phthalimide compound formed with R1 or R2, CH2, CH, and C, and

where at least one of R1 and R2 is substituted with a compounds having the chemical formula:

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$$O-(CH2)n-C-COOR7$$
R₆

where n is zero to five, where R_{δ} and R_{δ} are selected from the group consisting of hydrogen, halogen, substituted or unsubstituted C_{1-12} alkyl groups, carboxylic acid and ester groups, substituted or unsubstituted aromatic or heteroaromatic groups, and these moieties may be the same or different, or alkyl moieties of part of an aliphatic ring connecting R_{δ} and R_{δ} , and

where R_7 is a hydrogen, halogen, salt cation, metal, or substituted or unsubstituted $C_{1\text{-}6}$ alkyl group;

a compound having the formula:

where R_1 and R_2 each are a substituted or unsubstituted aromatic or heteroaromatic compound, or substituted or unsubstituted alkyl or heteroalkyl ring compound, or a substituted or unsubstituted phthalimide compound, and

where R1 and R2 may be the same or different,

where A is a chemical bridge which includes two to four chemical moieties bonded together between R_1 and R_2 ,

wherein said chemical moieties in A are selected from the group consisting of CO, O, S, SO_2 , NH, NR_3 where R_3 is a C_{1-6} alkyl group, NR_4 where R_4 includes two carbonyls as part of a phthalimide compound formed with R_1 or R_2 , CH₂, CH, and C, with the caveat that, except in the case where A contains two identical CH and C moieties positioned adjacent one another to form an alkene or alkyne, the chemical moieties in A are each different from one another, and

wherein at least one of R_1 or R_2 is substituted with a compound having the chemical formula:

$$O-(CH_2)_n$$
 $C-COOR_7$
 R_6

where n is zero to five,

where R_5 and R_6 are selected from the group consisting of hydrogen, halogen, substituted or unsubstituted C_{1-12} alkyl groups, carboxylic acid and ester, substituted or unsubstituted aromatic or heteroaromatic groups, and these moieties may be the same or different, or alkyl moieties of part of an aliphatic ring connecting R_5 and R_6 , and

where R_7 is a hydrogen, halogen, salt cation, metal, or substituted or unsubstituted $C_{1.6}$ alkyl group; and

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a compound having the formula:

$$\begin{matrix} \begin{matrix} \begin{matrix} H & O & H \\ & C & C \end{matrix} \\ R_1 - N - C - \begin{matrix} C & & & & & \\ & & & & & \\ & & & & & \\ \end{matrix} \end{matrix} \begin{matrix} \begin{matrix} R_2 & O \\ & C \\ & C \end{matrix} \\ R_3 \end{matrix}$$

where R_1 is selected from the group consisting of optionally substituted phenyl, adamantyl, napthyl, and indanyl, $R_{2\cdot 3}$ are alkyl moieties of a $C_{3\cdot 6}$ alkyl ring connecting R_2 and R_3 , and R_4 is a hydrogen, a monovalent salt cation, or a $C_{1\cdot 3}$ lower alkyl.

- The method of Claim 26, wherein the allosteric effector compound is 2-[4-(((3,5-dimethylanilino)carbonyl)methyl)phenoxy]-2-methylpropionic acid, or a physiologically acceptable salt thereof.
 - 29. The method of Claim 26, further comprising administering a chemotherapy agent.
 - 30. The method of Claim 29, wherein the chemotherapy agent is BCNU.